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Effects of Compounds on ^{125}I -GnRH-A Binding to hGnRH Receptors in HEK-293 Cell Membranes

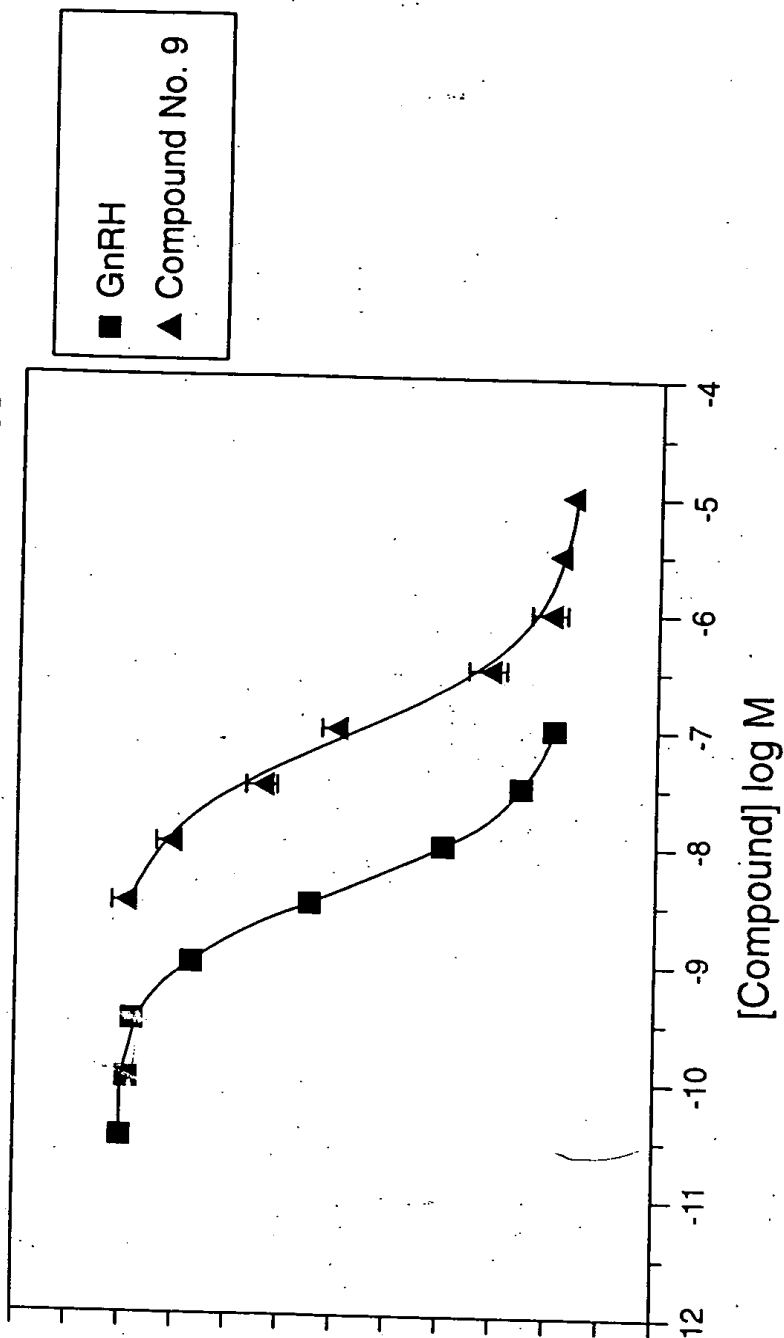


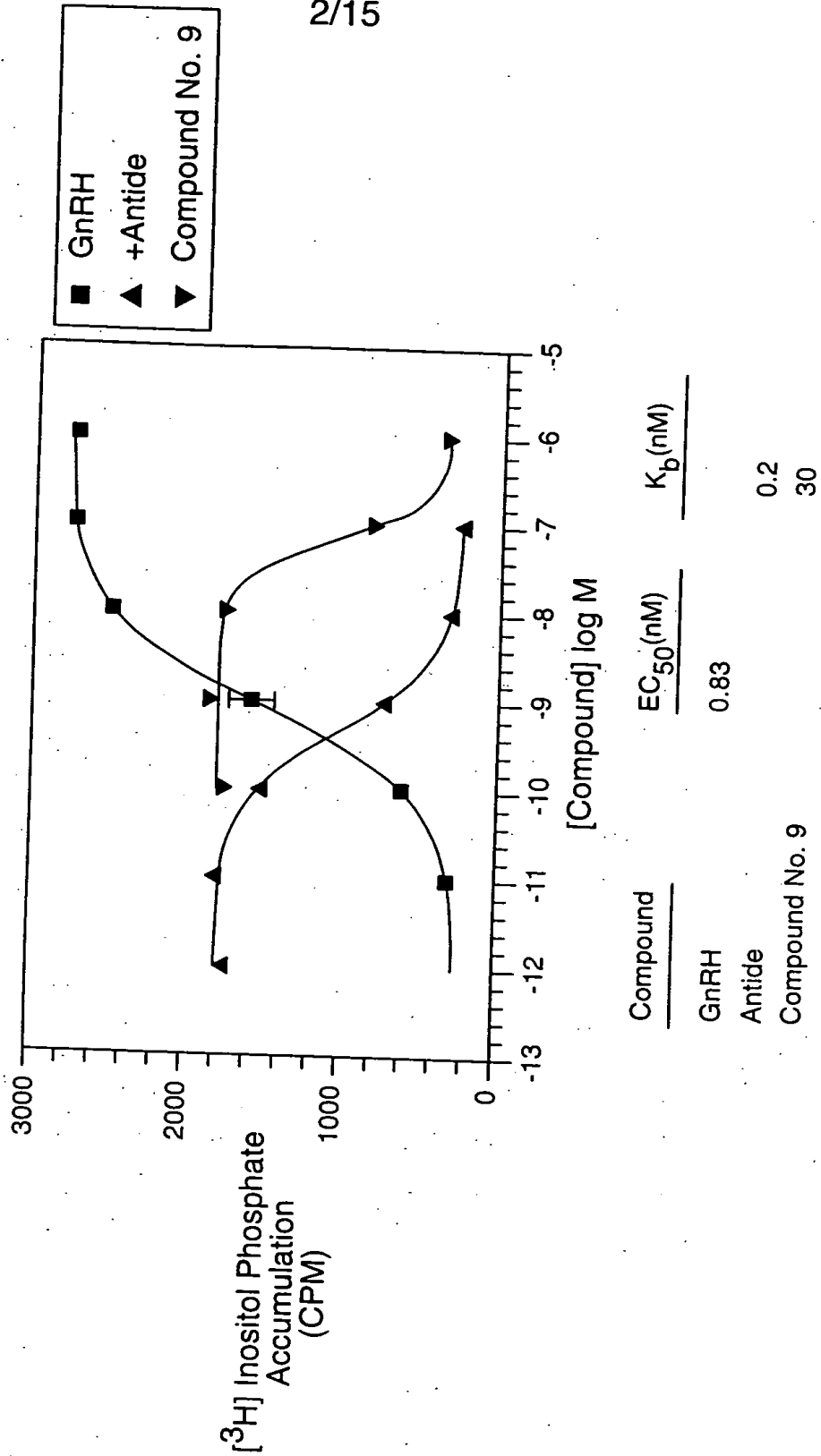
FIG. 1

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FIG. 2

Effects of Compounds on GnRH-stimulated (1 nM) total inositol phosphate accumulation in 293 cells expressing the hGnRH receptor



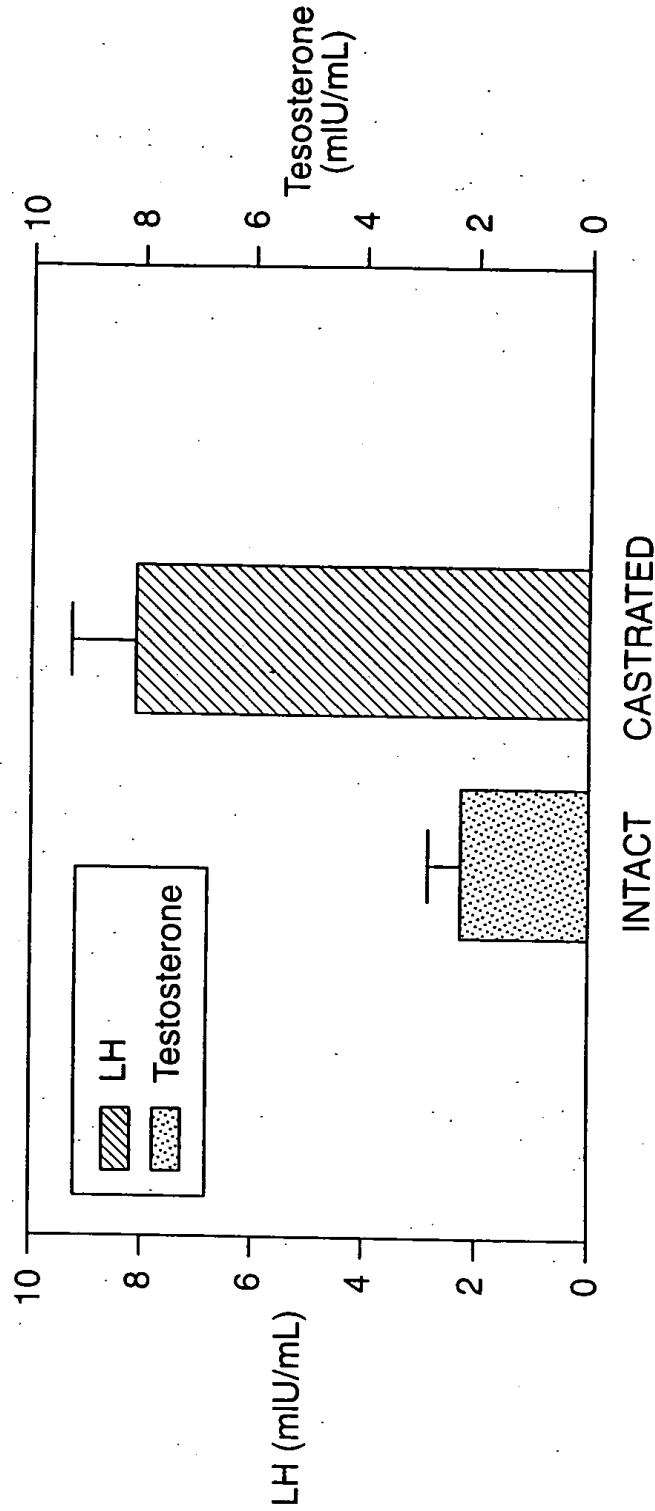
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FIG. 3

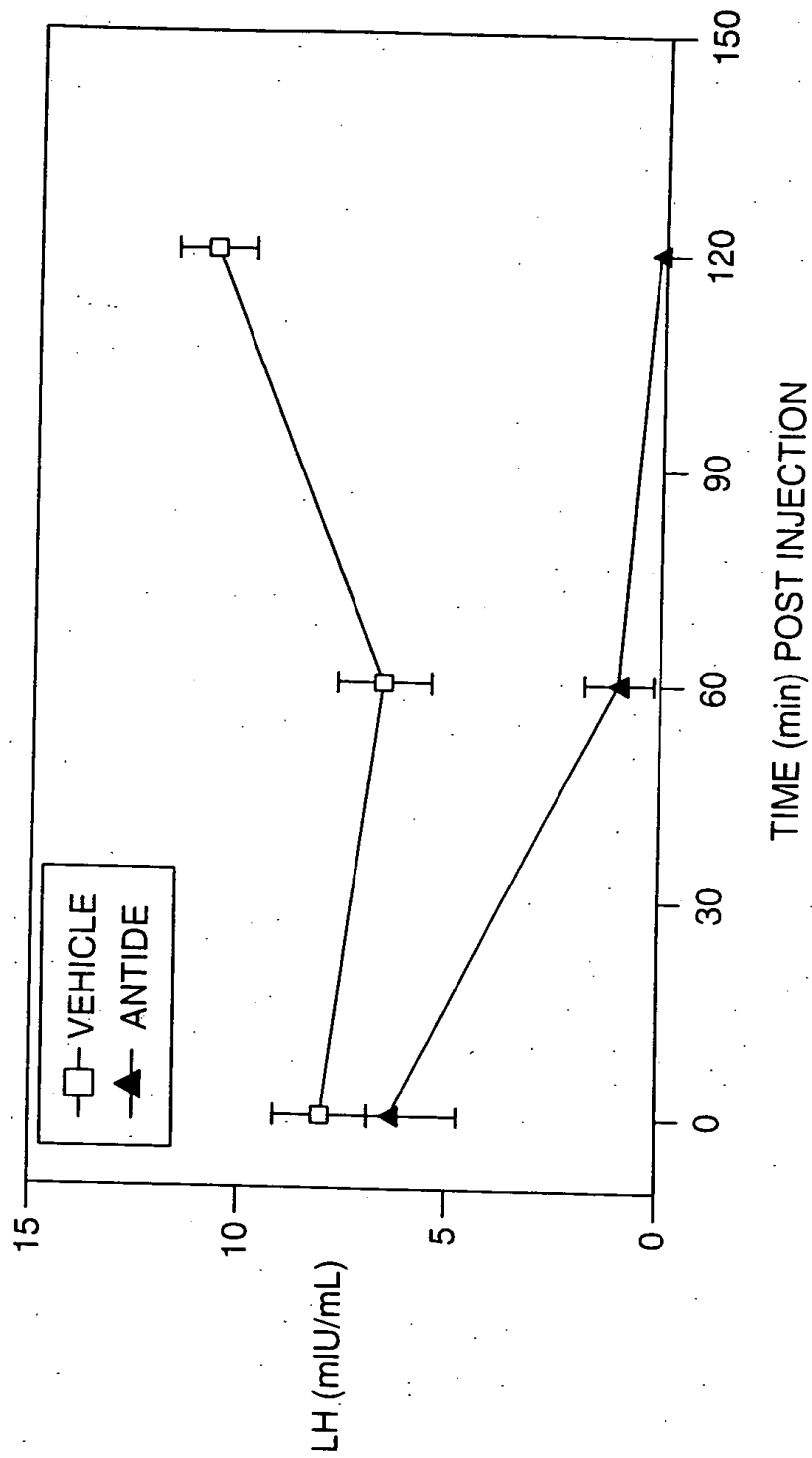
Characterization of a Castrated Rat Model
used to Evaluate GnRH Antagonists



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FIG. 4A

Antide (1.0 μ g;IV) suppresses LH levels
in the castrated rat model



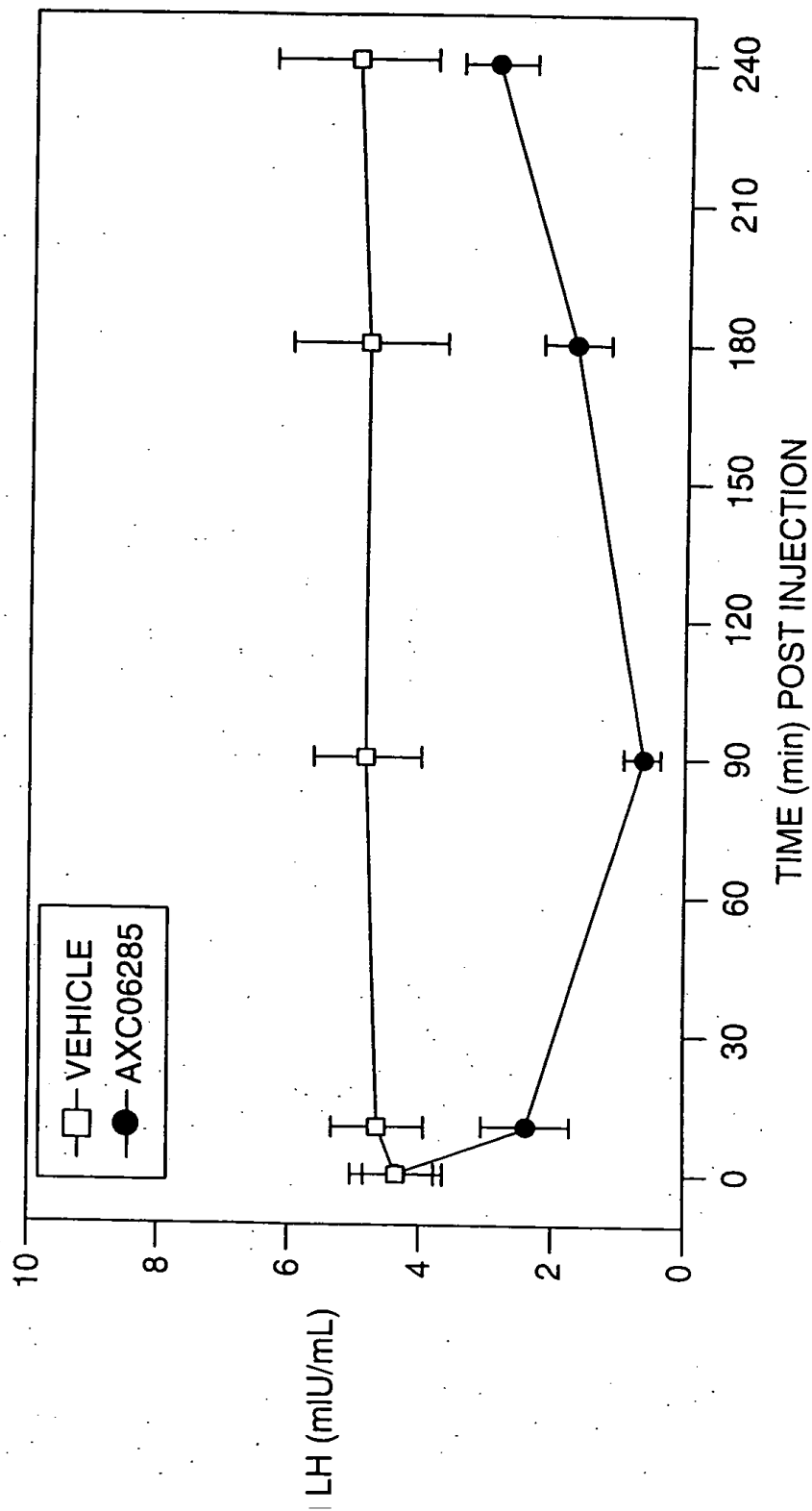
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Compound 11 (10mg/kg;IV) suppresses
LH levels in the castrated rat model

FIG. 4B



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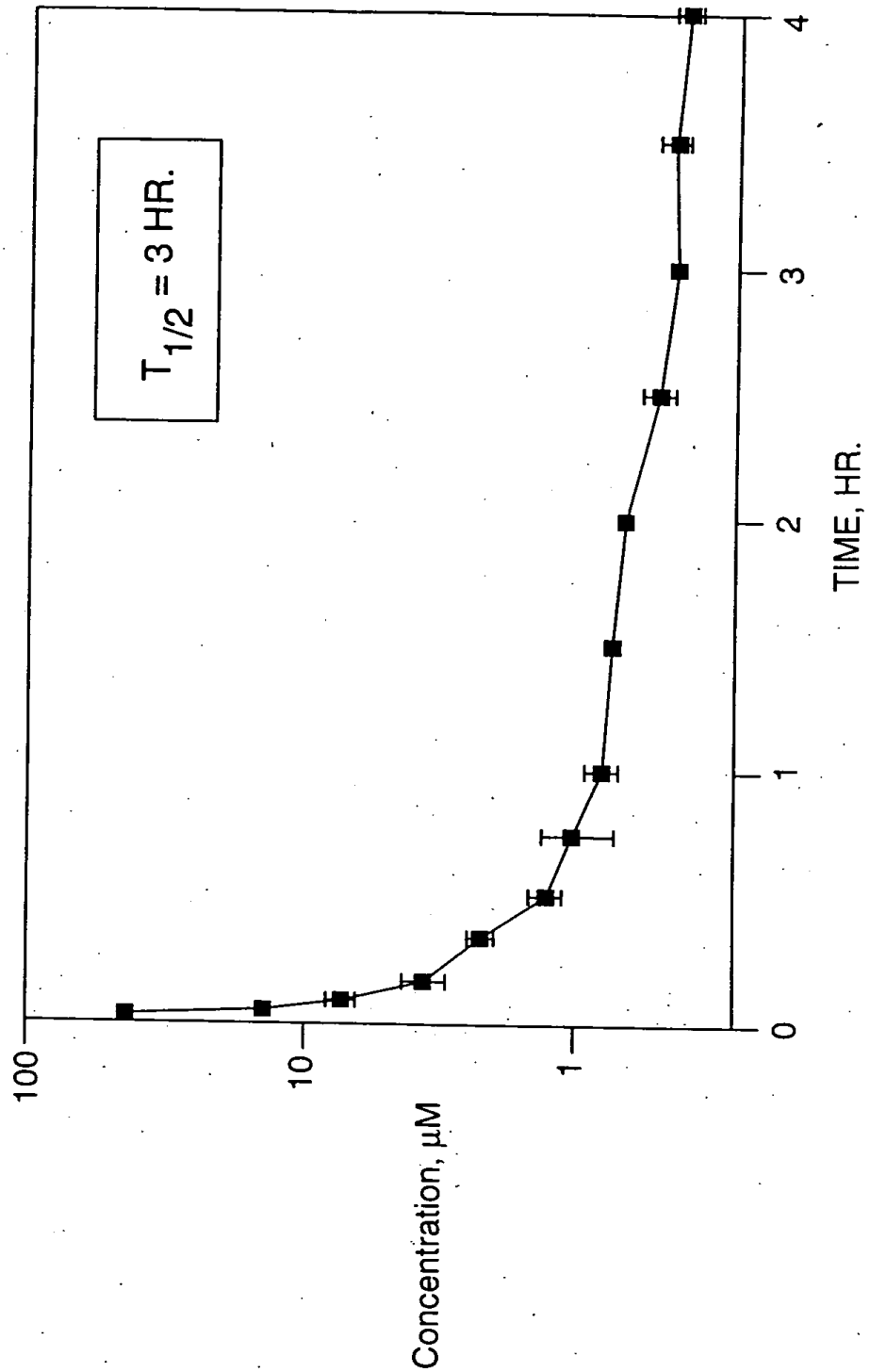
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FIG. 5

Compound 11 Dose - 10 mg/kg; n=3



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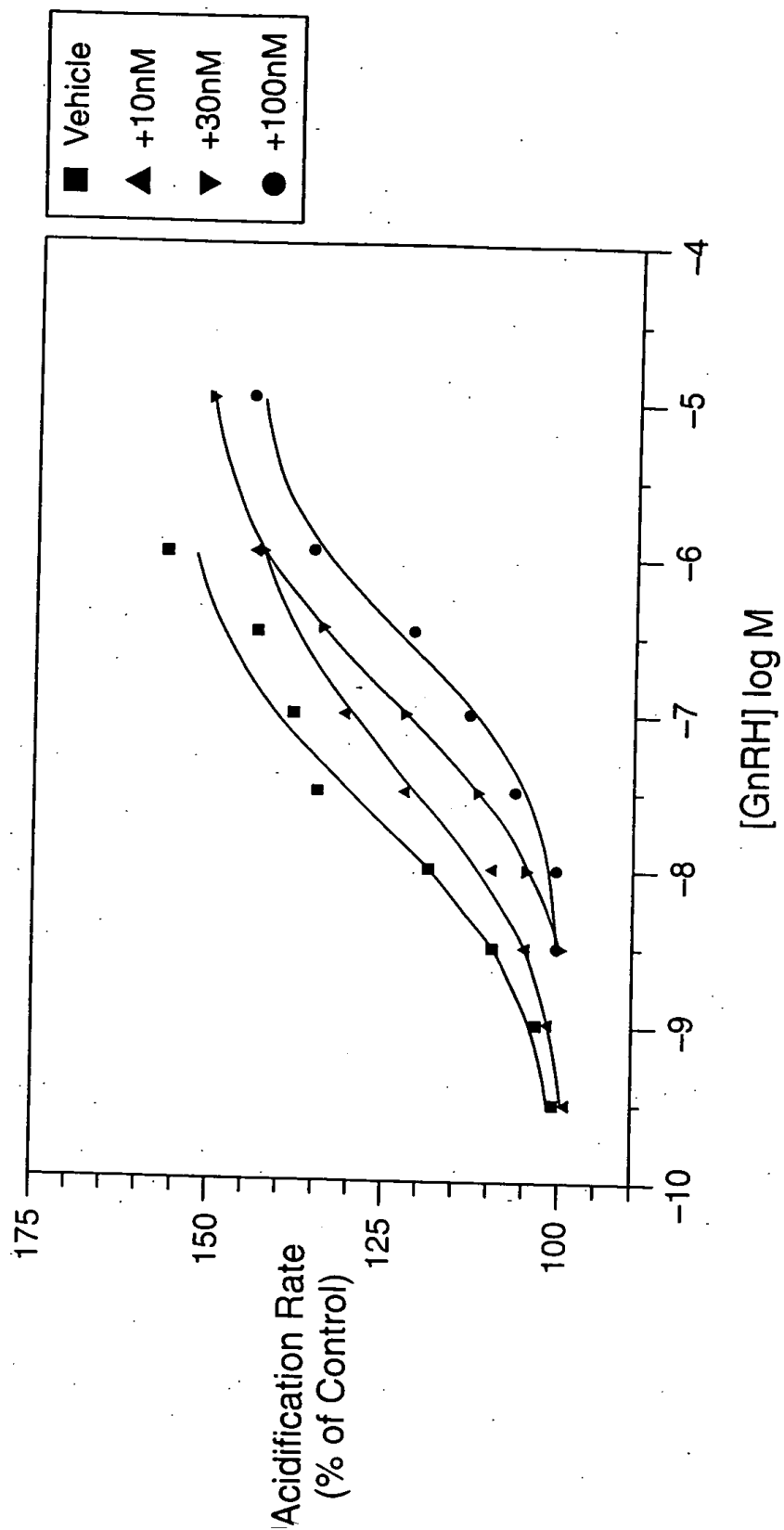


FIG. 6

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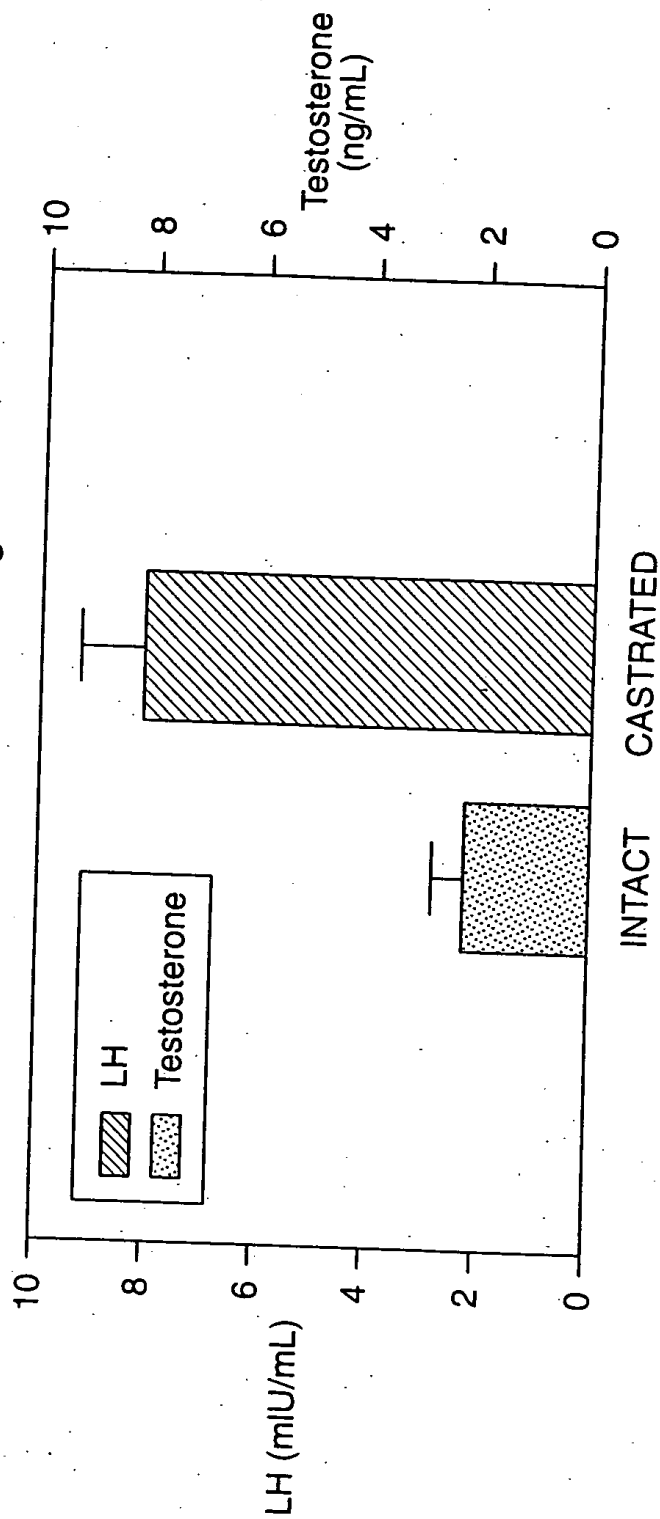


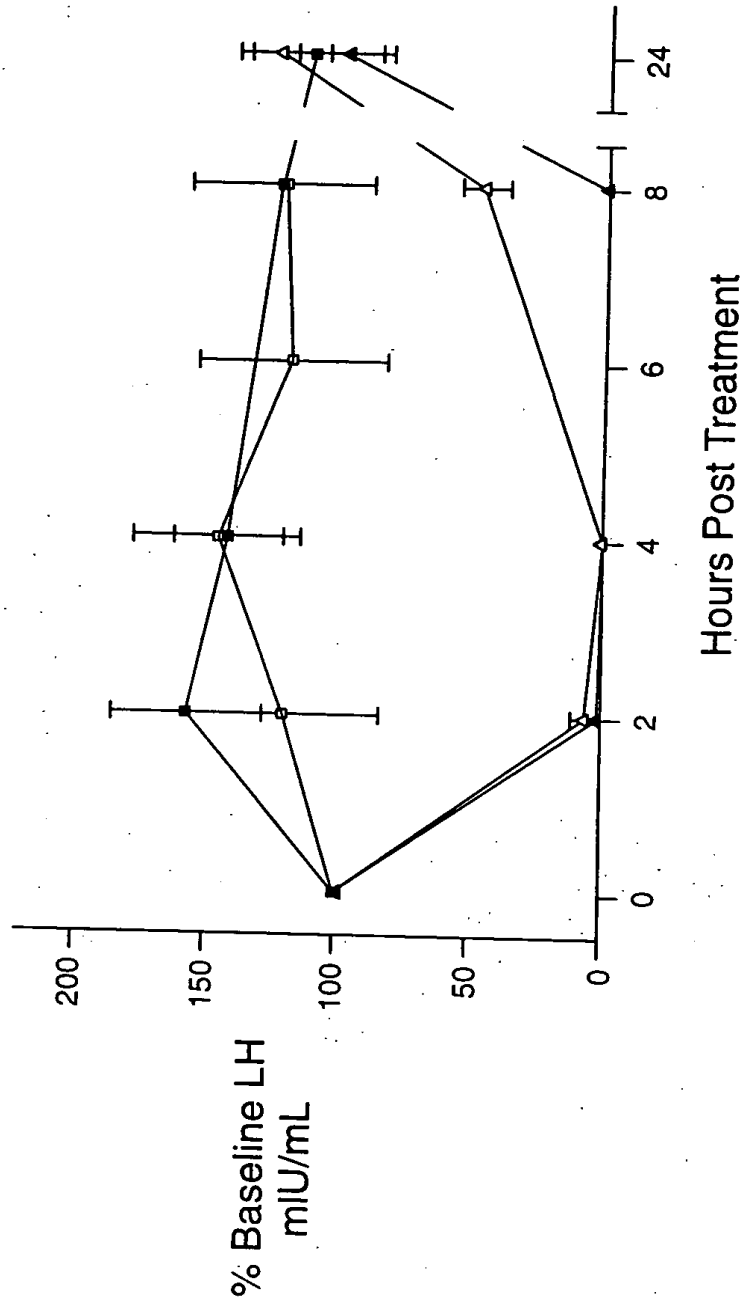
FIG. 7

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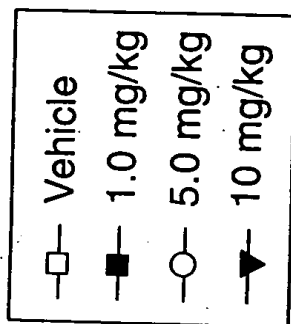
FIG. 8

Antide suppression of LH in
castrated rats is dose-dependent



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No. 134 (iv) suppression of LH is dose-dependent

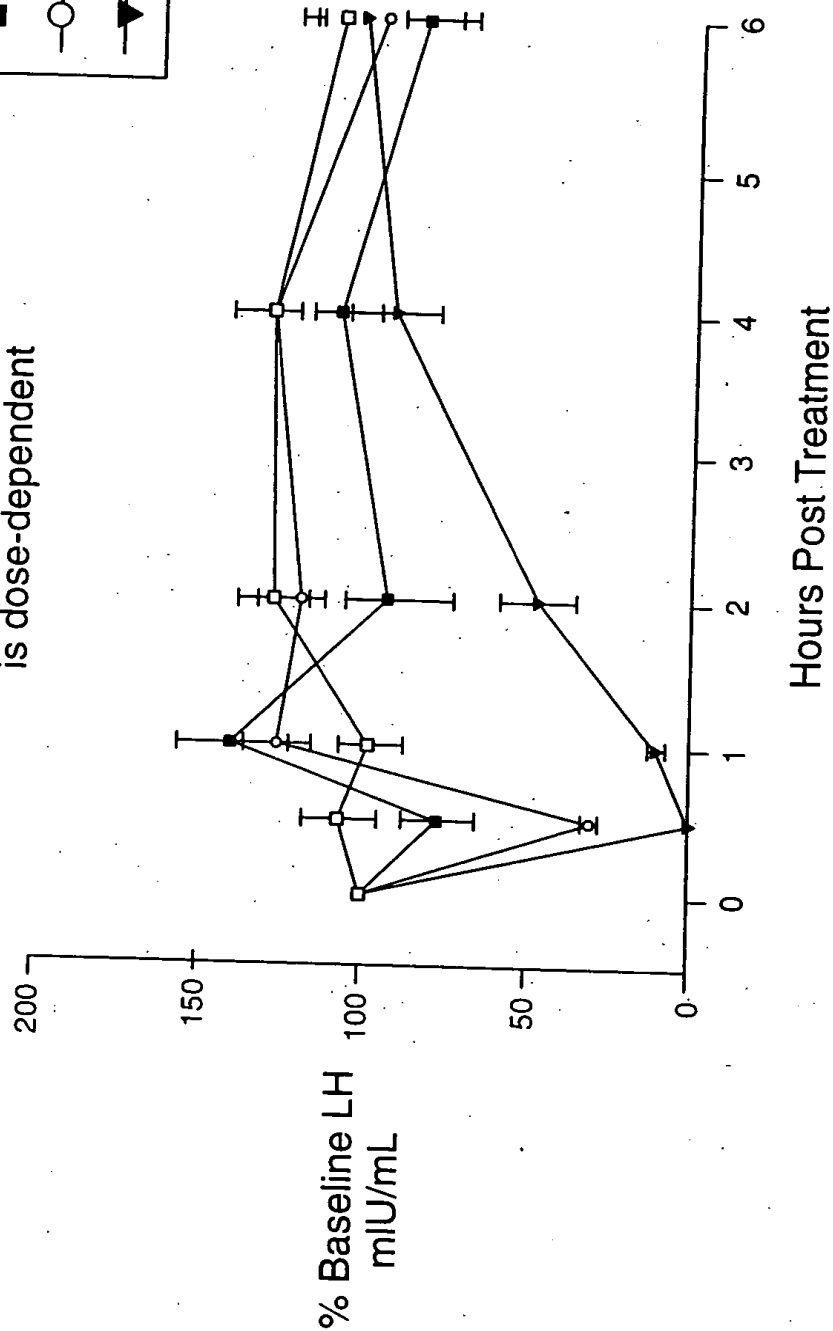
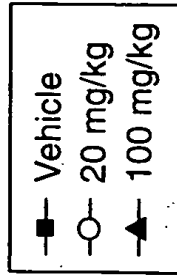


FIG. 9

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Oral delivery of No. 134 suppresses LH in castrated rats

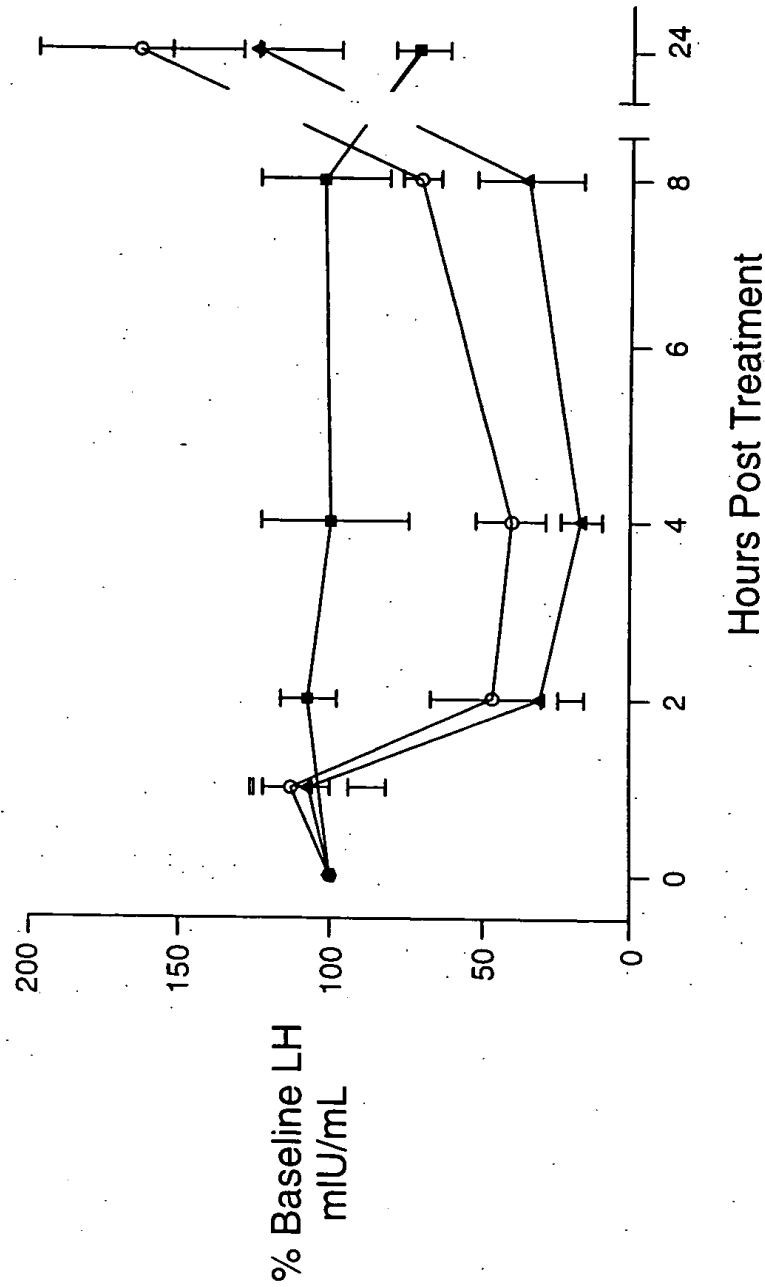


FIG. 10

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-□- Vehicle
 -▲- No. 134

IM delivery of No. 134 (20mg/kg)
suppresses LH in CX rats > 12 hrs.

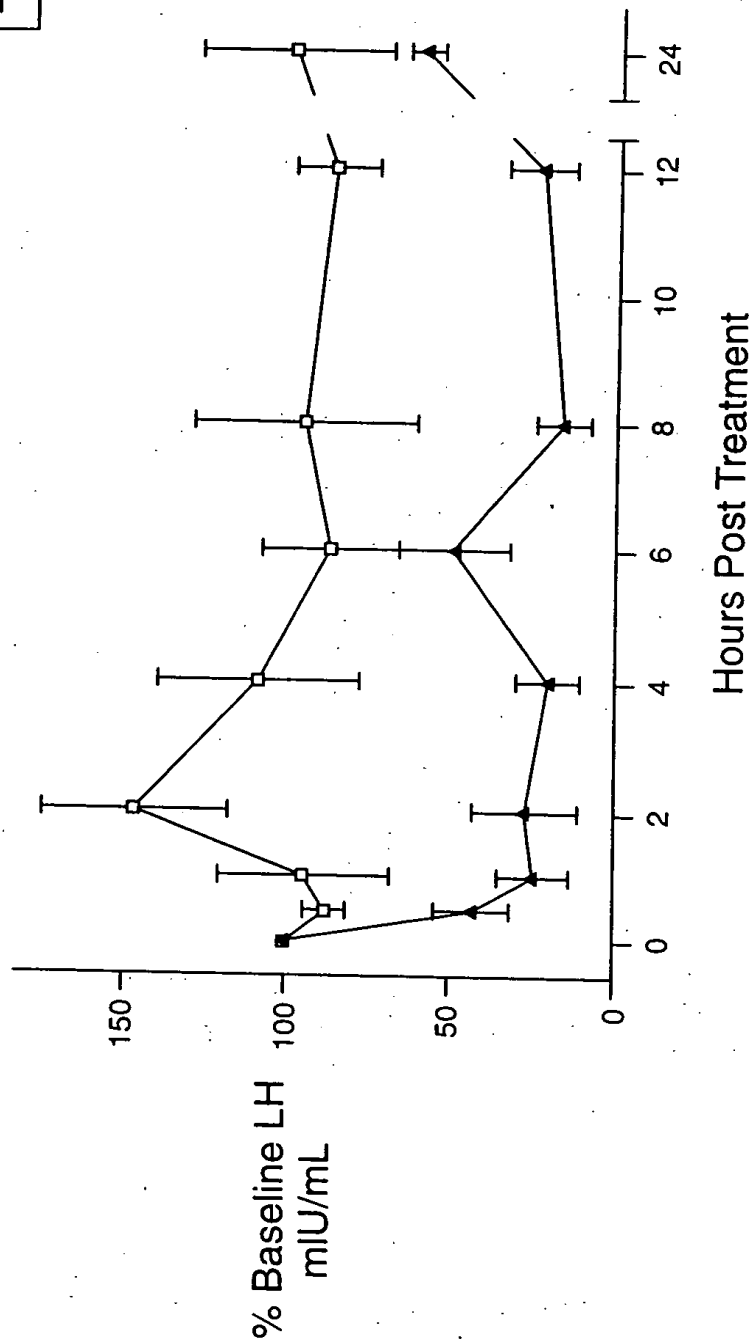


FIG. 11

FOUO 20 SEP 92 150

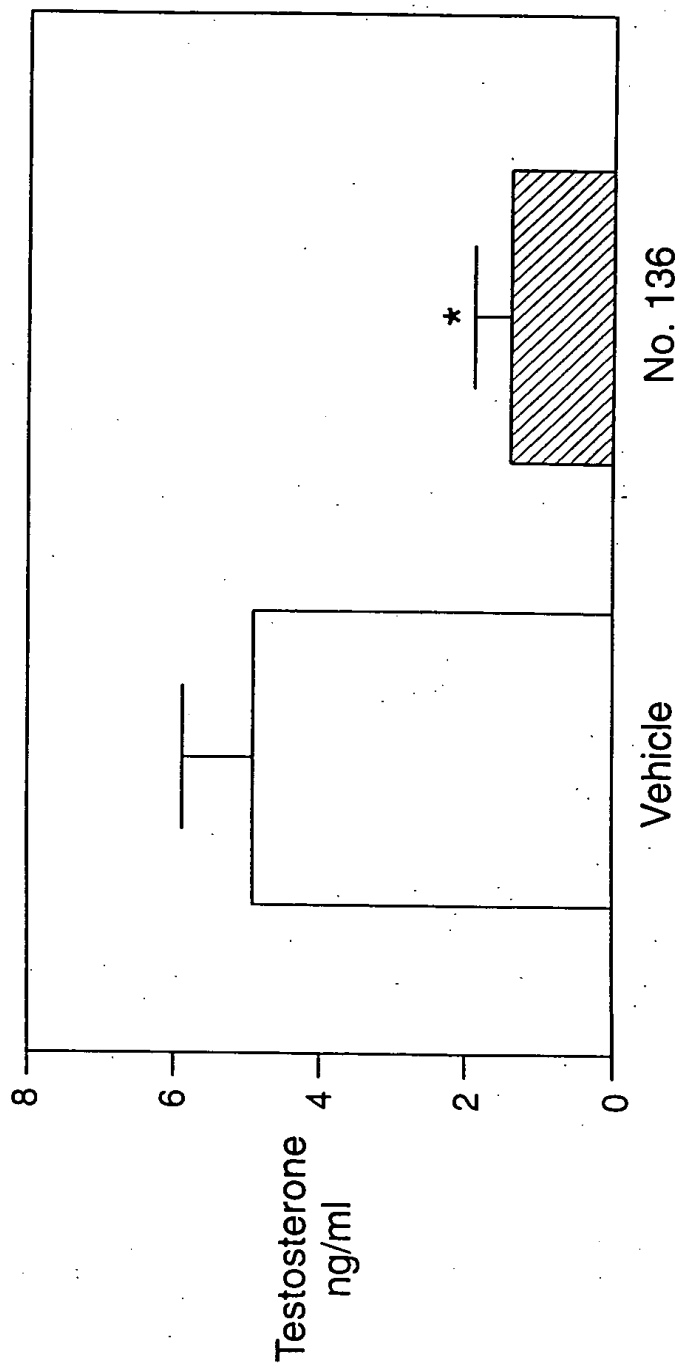
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FIG. 12

No. 136 (20 mg/kg;IP) suppresses testosterone 6 hours post treatment



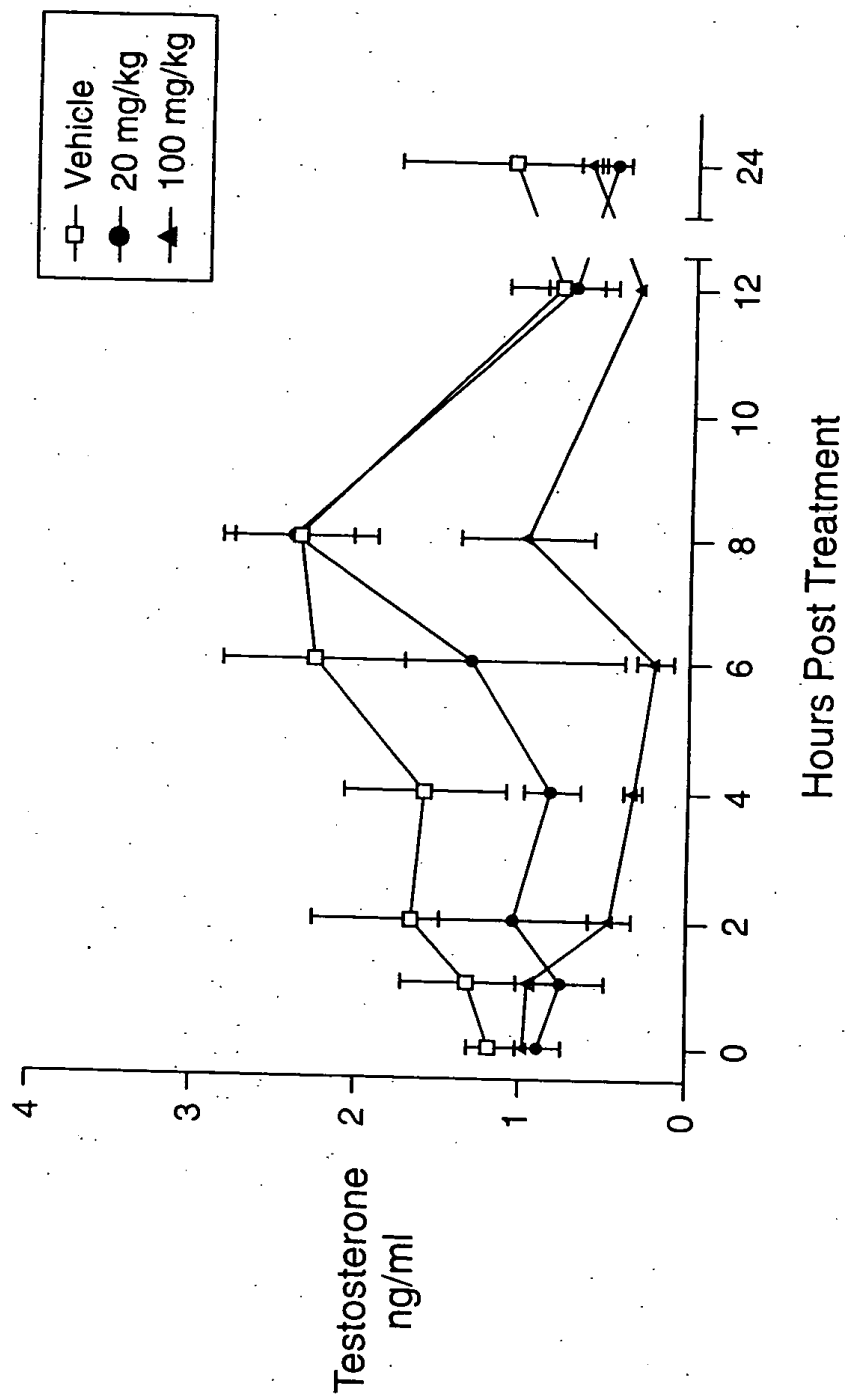
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FIG. 13

Effect of oral delivery of No. 134
on testosterone in intact rats



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FO0220" 9T2E9460

FIG. 14

Daily dosing of No. 134 (100 mg/kg;im)
suppresses testosterone in rats

